1. FENT COOPERATION TREAT?

From the INTERNATIONAL BUREAU **PCT** United States Patent and Trademark **NOTIFICATION OF ELECTION** (Box PCT) (PCT Rule 61.2) Crystal Plaza 2 Washington, DC 20231 **ETATS-UNIS D'AMERIQUE** Date of mailing (day/month/year) in its capacity as elected Office 16 December 1997 (16.12.97) Applicant's or agent's file reference International application No. SMW/BP5610415 PCT/GB97/01250 Priority date (day/month/year) International filing date (day/month/year) 08 May 1996 (08.05.96) 08 May 1997 (08.05.97) **Applicant** BALL, Kathryn, Lindsay et al 1. The designated Office is hereby notified of its election made: | X | in the demand filed with the International Preliminary Examining Authority on: 28 November 1997 (28.11.97) in a notice effecting later election filed with the International Bureau on: 2. The election was was not made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b).

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Authorized officer

R. Raissi

Telephone No.: (41-22) 338.83.38

Facsimile No.: (41-22) 740.14.35

wie ~

PATENT COOPERATION TREATY

,~			
	From the INTERNATIONAL BUREAU		
PCT	То:		
NOTIFICATION OF THE RECORDING OF A CHANGE (PCT Rule 92bis.1 and Administrative Instructions, Section 422) Date of mailing (day/month/year)	WALTON, Seán, M. Mewburn Ellis York House 23 Kingsway London WC2B 6HP ROYAUME-UNI		
20 August 1997 (20.08.97)			
Applicant's or agent's file reference SMW/BP5610415	IMPORTANT NOTIFICATION		
International application No. PCT/GB97/01250	International filing date (day/month/year) 08 May 1997 (08.05.97)		
The following indications appeared on record concerning:			
X the applicant the inventor	the agent the common representative		
Name and Address	State of Nationality State of Residence GB GB		
UNIVERSITY OF DUNDEE Dundee	Telephone No.		
DD1 4HN-UNI United kingdon			
	Facsimile No.		
	Teleprinter No.		
2. The International Bureau hereby notifies the applicant that th	e following change has been recorded concerning:		
X the person the name the add			
Name and Address	State of Nationality State of Residence		
CYCLACEL LIMITED. Marquis House	Telephone No.		
67-68 Jermyn Street London SW1Y 6NY			
United kingdom	Facsimile No.		
·	Teleprinter No.		
Further observations, if necessary: Please note that a new Power of Attorney is needed. Also, pland residence of the new applicant.	lease provide the Nationality		
4. A copy of this notification has been sent to:			
X the receiving Office	X the designated Offices concerned		
X the International Searching Authority	the elected Offices concerned		
the International Preliminary Examining Authority	other:		
	Authorized officer		
The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	I. Britel		

Telephone No.: (41-22) 338.83.38



PCT

NOTICE INFORMING THE APPLICANT OF THE COMMUNICATION OF THE INTERNATIONAL APPLICATION TO THE DESIGNATED OFFICES

(PCT Rule 47.1(c), first sentence)

From the INTERNATIONAL BUREAU

Į To

WALTON, Seán, M. Mewburn Ellis York House 23 Kingsway London WC2B 6HP ROYAUME-UNI

RECEIVED
2 1 NOV 1997

Date of mailing (day/month/year) .

13 November 1997 (13.11.97)

Applicant's or agent's file reference

SMW/BP5610415
International application No.

PCT/GB97/01250

International filing date (day/month/year) 08 May 1997 (08.05.97)

Priority date (day/month/year) 08 May 1996 (08.05.96)

IMPORTANT NOTICE

Applicant

CYCLACEL LIMITED et al

 Notice is hereby given that the International Bureau has communicated, as provided in Article 20, the international application to the following designated Offices on the date indicated above as the date of mailing of this Notice: AU,BR,CA,CN,EP,IL,JP,KP,KR,NO,PL,SK,US

In accordance with Rule 47.1(c), third sentence, those Offices will accept the present Notice as conclusive evidence that the communication of the international application has duly taken place on the date of mailing indicated above and no copy of the international application is required to be furnished by the applicant to the designated Office(s).

2. The following designated Offices have waived the requirement for such a communication at this time:

AL,AM,AP,AT,AZ,BA,BB,BG,BY,CH,CU,CZ,DE,DK,EA,EE,ES,FI,GB,GE,GH,HU,IS,KE,KG,KZ,LC,LK,LR,LS,LT,LU,LV,MD,MG,MK,MN,MW,MX,NZ,OA,PT,RO,RU,SD,SE,SG,SI,TJ,TM,TR,TT,UA,UG,UZ,VN,YU

The communication will be made to those Offices only upon their request. Furthermore, those Offices do not require the applicant to furnish a copy of the international application (Rule 49.1(a-bis)).

 Enclosed with this Notice is a copy of the international application as published by the International Bureau on 13 November 1997 (13.11.97) under No. WO 97/42222

REMINDER REGARDING CHAPTER II (Article 31(2)(a) and Rule 54.2)

If the applicant wishes to postpone entry into the national phase until 30 months (or later in some Offices) from the priority date, a demand for international preliminary examination must be filed with the competent International Preliminary Examining Authority before the expiration of 19 months from the priority date.

It is the applicant's sole responsibility to monitor the 19-month time limit.

Note that only an applicant who is a national or resident of a PCT Contracting State which is bound by Chapter II has the right to file a demand for international preliminary examination.

REMINDER REGARDING ENTRY INTO THE NATIONAL PHASE (Article 22 or 39(1))

If the applicant wishes to proceed with the international application in the national phase, he must, within 20 months or 30 months, or later in some Offices, perform the acts referred to therein before each designated or elected Office.

For further important information on the time limits and acts to be performed for entering the national phase, see the Annex to Form PCT/IB/301 (Notification of Receipt of Record Copy) and Volume II of the PCT Applicant's Guide.

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Authorized officer

J. Zahra

Telephone No. (41-22) 338.83.38

Facsimile No. (41-22) 740.14.35



NOTICE INFORMING THE APPLICANT OF THE COMMUNICATION OF THE INTERNATIONAL APPLICATION TO THE DESIGNATED OFFICES

Date of mailing (day/month/year) 13 November 1997 (13.11.97)	IMPORTANT NOTICE		
Applicant's or agent's file reference SMW/BP5610415	International application No. PCT/GB97/01250		

The applicant is hereby notified that, at the time of establishment of this Notice, the time limit under Rule 46.1 for making amendments under Article 19 has not yet expired and the International Bureau had received neither such amendments nor a declaration that the applicant does not wish to make amendments.



RECORDS NO EO.

NACE OF PRIORITY DOCUMENTS

(PCT Administrative Instructions, Section 411)

From the INTERNATIONAL BUREAU

To:

WALTON, Seán, M. Mewburn Ellis York House 23 Kingsway London WC2B 6HP ROYAUME-UNI 1 2 JUN 1997

Date of mailing (day/month/year)

04 June 1997 (04.06.97)

Applicant's or agent's file reference

\$MW/BP5610415

IMPORTANT NOTIFICATION

International application No. PCT/GB97/01250

International filing date (day/month/year) 08 May 1997 (08.05.97)

Priority date (day/month/year) 08 May 1996 (08.05.96)

Applicant

UNIVERSITY OF DUNDEE et al

The applicant is hereby notified of the date of receipt by the International Bureau of the priority document(s) relating to the following application(s):

Priority application No:

Priority date:

Priority country:

Date of receipt of priority document:

9609521.1

08 May 1996 (08.05.96)

GB

02 Jun 1997 (02.06.97)

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland

Authorized officer

州CG-M.C. Guillemot

Facsimile No.: (41-22) 740.14.35

Telephone No.: (41-22) 338.83.38

From the INTERNATIONAL BUREAU

PCT

NOTIFICATION CONCERNING SUBMISSION OF PRIORITY DOCUMENTS

(PCT Administrative Instructions, Section 411)

WALTON, Seán, M. Mewburn Ellis York House 23 Kingsway London WC2B 6HP

1 9 JUN 1997

ROYAUME-UNI

Date of mailing (day/month/year)
13 June 1997 (13.06.97)

Applicant's or agent's file reference

International application No.

PCT/GB97/01250

International filing date (day/month/year)
08 May 1997 (08.05.97)

Priority date (day/month/year) 08 May 1996 (08.05.96)

IMPORTANT NOTIFICATION

Applicant

UNIVERSITY OF DUNDEE et al

The applicant is hereby notified of the date of receipt by the International Bureau of the priority document(s) relating to the following application(s):

Priority application No:

Priority date:

Priority country:

Date of receipt of priority document:

9621314.5

09 Oct 1996 (09.10.96)

GB

10 Jun 1997 (10.06.97

RECORDS NOTED

COMPUTES ENT'D

REGISTER ENT'S

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THARY ENT'D

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland

Facsimile No.: (41-22) 740.14.35

Authorized officer

りん Guillemot

Telephone No.: (41-22) 338.83.38

ATENT COOPERATION TREAT From the INTERNATIONAL BUREAU ARY ENT D NOTIFICATION OF THE RECORDING WALTON, Seán, M. OF A CHANGE Mewburn Ellis York House (PCT Rule 92bis.1 and 23 Kingsway Administrative Instructions, Section 422) London WC2B 6HP **ROYAUME-UNI** Date of mailing (day/month/year) 20 August 1397 (20.08.97 Applicant's or agent's file reference IMPORTANT NOTIFICATION SMW/BP\$610415 International filing date (day/month/year) International application No. PCT/GB97/01250 08 May 1997 (08.05.97) 1. The following indications appeared on record concerning: X the applicant the inventor the agent the common representative State of Nationality State of Residence Name and Address GB UNIVERSITY OF DUNDEE Dundee Telephone No. DD1 4HN-UNI United kingdon Facsimile No. Teleprinter No. 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning: the residence X the person the address the nationality the name State of Residence State of Nationality Name and Address CYCLACEL LIMITED Marquis House Telephone No. 67-68 Jermyn Street London SW1Y 6NY United kingdom Facsimile No. Teleprinter No. 3. Further observations, if necessary: Please note that a new Power of Attorney is needed. Also, please provide the Nationality and residence of the new applicant. 4. A copy of this notification has been sent to: the designated Offices concerned

The Internati nal Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland

the International Preliminary Examining Authority

the International Searching Authority

Authorized officer

I. Britel

the elected Offices concerned

Facsimile No.: (41-22) 740.14.35

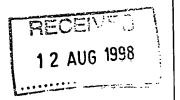
the receiving Office

other:



INTEH TATIONAL PRELIMINARY EXAMINING AUTHORITY

KIDDLE, Simon J. MEWBURN ELLIS York House 23 Kingsway London WC2B 6HP GRANDE BRETAGNE



NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL PRELIMINARY **EXAMINATION REPORT**

(PCT Rule 71.1)

Date of mailing (day/month/year)

10.08.98

Applicant's or agent's file reference SJK/BP5610415

International application No. PCT/GB97/01250

International filing date (day/month/year) 08/05/1997

Priority date (day/month/year) 08/05/1996

IMPORTANT NOTIFICATION

Applicant

CYCLACEL LIMITED et al.

- 1. The applicant is hereby notified that this International Preliminary Examining Authority transmits herewith the international preliminary examination report and its annexes, if any, established on the international application.
- 2. A copy of the report and its annexes, if any, is being transmitted to the International Bureau for communication to all the elected Offices.
- 3. Where required by any of the elected Offices, the International Bureau will prepare an English translation of the report (but not of any annexes) and will transmit such translation to those Offices.

4. REMINDER

The applicant must enter the national phase before each elected Office by performing certain acts (filing translations and paying national fees) within 30 months from the priority date (or later in some Offices) (Article 39(1)) (see also the reminder sent by the International Bureau with Form PCT/IB/301).

Where a translation of the international application must be furnished to an elected Office, that translation must contain a translation of any annexes to the international preliminary examination report. It is the applicant's responsibility to prepare and furnish such translation directly to each elected Office concerned.

For further details on the applicable time limits and requirements of the elected Offices, see Volume II of the PCT Applicant's Guide.

Name and mailing address of the IPEA/

Authorized officer

European Patent Office D-80298 Munich

Heisel, M

Tel. (+49-89) 2399-0. Tx: 523656 epmu d

Tel. (+49-89) 2399-8051

Fax: (+49-89) 2399-4465

PATENT COOPERATION TREATY

PCT

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

REC'D

1 3 AUG 1998

(PCT Article 36 and Rule 70)

Applicant's or	agent	s file reference	FOR FURTHER ACTION	See	Notification of Transmittal of International
SJK/BP56	SJK/BP5610415 FOR FURTHER ACTION Preliminary Examination Report (PCT/IPEA/416				iminary examination Report (PC1/IPEA/416)
International	applica	tion No.	International filing date (day/month/year	-)	Priority date (day/month/year)
PCT/GB97	/0125	50	08/05/1997		08/05/1996
International	Patent	Classification (IPC) or na	ational classification and IPC		
C07K14/47	7				
					*
Applicant					
CYCLACE	L LIM	IITED et al.			
1. This int	ernati	onal preliminary exam	nination report has been prepared by	this In	ternational Preliminary Examining Authority
and is t	ransm	nitted to the applicant	according to Article 36.		
2. This Ri	EPOR	T consists of a total of	f 6 sheets, including this cover she	∋t.	
57 74	.: _	art is also secompani	ed by ANNEXES, i.e., sheets of the	descrip	ition, claims and/or drawings
	خا حاجت	ava baan amandad ar	nd are the basis for this report and/o	r sneet	s containing rectifications made
be	fore t	his Authority (see Rule	e 70.16 and Section 607 of the Adm	nıstratı	ve instructions under the PC1).
			f 10 shoots		
These	annex	es consist of a total o	i io sileets.		
3. This re	port c	ontains indications rel	ating to the following items:		
I ⊠ Basis of the report					
1 1	Ø	Priority			
1 111	Ø		of opinion with regard to novelty, inv	entive s	step and industrial applicability
IV		Lack of unity of inve			
V	_ ⊠	Reasoned statemen	t under Article 35(2) with regard to r	ovelty,	inventive step or industrial applicability;
		citations and explan	ations supporting such statement		·
VI		Certain documents			
VII			e international application		
VIII	\boxtimes	Certain observation	s on the international application		
Date of sub	mis sio	n of the demand	Date of co	mpletior	of this report
		,			1 0. 08. 98
28/11/19	97				
Name	nailin-	address of the IPFA/	Authorized	officer	

European Patent Office D-80298 Munich

Name and mailing address of the IPEA/

Tel. (+49-89) 2399-0. Tx: 523656 epmu d

Fax: (+49-89) 2399-4465

Stolz. B

Telephone No. (+49-89) 2399-8416



INTERNATIONAL PRELIMINARY **EXAMINATION REPORT**

International application No. PCT/GB97/01250

I. E	3as	is	of	the	r	port
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1. This report has been drawn on the basis of (substitute sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to the report since they do not contain amendments.): Description, pages: as originally filed 1-78 Claims, No.: 24/07/1998 24/07/1998 with letter of 1-36 as received on Drawings, sheets: 1/10-10/10 as originally filed 2. The amendments have resulted in the cancellation of: ☐ the description, pages: the claims, Nos.: the drawings, sheets: 3. This report has been established as if (some of) the amendments had not been made, since they have been considered to go beyond the disclosure as filed (Rule 70.2(c)): 4. Additional observations, if necessary: II. Priority 1.

This report has been established as if no priority had been claimed due to the failure to furnish within the prescribed time limit the requested: copy of the earlier application whose priority has been claimed. translation of the earlier application whose priority has been claimed.

2. This report has been established as if no priority had been claimed due to the fact that the priority claim has

been found invalid.



INTERNATIONAL PRELIMINARY **EXAMINATION REPORT**

International application No. PCT/GB97/01250

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The	us fo	r the purposes of this report, the international filing date indicated above is considered to be the relevant date
3.	Addi	itional observations, if necessary:
	see	separate sheet
11.	Non	establishment of opinion with regard to novelty, inventive step and industrial applicability
Γh or	e que to be	estions whether the claimed invention appears to be novel, to involve an inventive step (to be non-obvious). industrially applicable have not been examined in respect of:
		the entire international application.
	×	claims Nos. 24-28, 34-36.
рe	caus	
	Ø	the said international application, or the said claims Nos. 24-28, 34-36 relate to the following subject matter which does not require an international preliminary examination (<i>specify</i>):
		see separate sheet
		the description, claims or drawings (indicate particular elements below) or said claims Nos. are so unclear that no meaningful opinion could be formed (specify):
		the claims, or said claims Nos. are so inadequately supported by the description that no meaningful opinion could be formed.
		no international search report has been established for the said claims Nos

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/GB97/01250

- V. Reasoned statem int under Articl 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement
- 1. Statement

Novelty (N)

Yes: No: Claims 1-23, 29-33

Claims

Inventive step (IS)

Claims

ns 1-23, 29-33

Yes: No:

o: Claims

Industrial applicability (IA)

Yes:

Claims 1-23, 29-33

No:

: Claims

2. Citations and explanations

see separate sheet

VIII. Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made:

see separate sheet

EXAMINATION REPORT - SEPARATE SHEET

1. Priority

Priority documents have not been available at the time of establishing this report. It has however been established under the assumption of valid priority rights. Should this not hold true, the following documents, cited in the ISR as P-documents, might become important: Ball et al., Current Biology, 1997, vol. 7, no. 1, pp. 71-80; and Chen et al., Mol. Cell. Biol., 1996, vol. 16, no. 9, pp. 4673-4682.

2. No opinion

The subject matter of claims 24 to 28, and 34 to 36 relates to methods for the treatment of the human or animal body. In accordance with Rule 67.1 PCT, an opinion for these claims is not established.

3. Reasoned statement

3.1. Basically, the application discloses regions or segments of p21 important for interaction with cyclin D1 and/or Cdk4. Claimed are (i) methods of identifying compounds interfering with the interaction between p21 and cyclin D1 and/or Cdk4 by using a set of peptides including defined fragments, (ii) the use of those peptides in such screening methods, (iii) the use of p21 fragments or derivatives in the manufacture of medicaments or for medical treatment of cdk4 mediated disease, and (iv) the use of defined p21 fragments in order to interfere with the interaction between p21 and cyclin D1 and/or Cdk4.

3.2. Novelty (Art. 33(2) PCT)

Assays to identify compounds affecting the interaction of p21 with cyclin D1 and/or Cdk4 by using fragments of p21 have not been disclosed in the prior art. Claims 1 to 23 are thus novel. The various uses of p21 fragments of claims 24 to 33 have also not been disclosed in the available prior art.

EXAMINATION REPORT - SEPARATE SHEET

3.3. Inventive step (Art. 33(3) PCT)

The prior art contains a number of studies to identify domains of p21 important for its interaction with various binding partners. But it is silent with respect to domains important for interaction with cyclin D1 and cdk4. This information cannot be derived from the prior art in an obvious manner. Therefore, also the use of such domains (or fragments) in methods according to claims 1 to 23, and claims 31 to 33, and the defined peptide of claims 29 and 30 cannot be derived from the prior art.

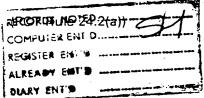
Certain observations 4.

- 4.1. Claim 1 as presently worded could also include the use of full-length p21 because p21 could be described as "a substance which includes" one or several of the listed fragments. The prior art describes studies of p21 and mutants thereof with respect to their ability to interact with cyclin D1 and to their ability to affect activity of the cyclin D1/Cdk4 complex. The use of this system to search for interfering compounds is obvious to those of skill.
- 4.2. The term "derivative or analogue" used throughout the claims is vague and renders the scope of the claims unclear.
- 4.3. On several occasions "functional mimetics" are mentioned in the claims without any technical specification. The function which is to be mimicked is not specified, and the application does not provide a single example. Thus, the scope of those claims cannot be unambiguously established. Moreover, even if the function was defined, inventive step would be required to find mimetics of any of the defined peptides.
- 4.4. Claim 9 refers to "a p21 mediated effect on cdk4 activity" without specifying what effect. It seems thus impossible to determine when a method is falling within the scope of claim 9.

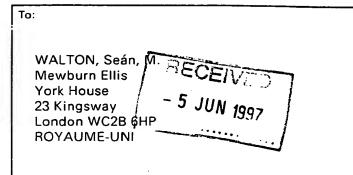
The same argument seems to apply to the "cdk4 activity" of claim 14, and "the property of inhibiting cdk4" of claim 29.

NOTIFICATION OF RECEIPT OF RECORD COPY

PCT



From the INTERNATIONAL BUREAU



Date of mailing (day/month/year) 26 May 1997 (26.05.97)	IMPORTANT NOTIFICATION
Applicant's or agent's file reference. SMW/BP5610415	International application No. PCT/GB97/01250

The applicant is hereby notified that the International Bureau has received the record copy of the international application as detailed below.

Name(s) of the applicant(s) and State(s) for which they are applicants:

UNIVERSITY OF DUNDEE (for all designated States except US) BALL, Kathryn, Lindsay et al (for US)

International filing date Priority date(s) claimed 08 May 1997 (08.05.97)

08 May 1996 (08.05.96) 09 October 1996 (09.10.96)

Date of receipt of the record copy by the International Sureau

26 May 1997 (26.05.97)

List of designated Offices

AP:GH,KE,LS,MW,SD,SZ,UG

EA:AM,AZ,BY,KG,KZ,MD,RU,TJ,TM

EP:AT,BE,CH,DE,DK,ES,FI,FR,GB,GR,IE,IT,LU,MC,NL,PT,SE

OA:BF,BJ,CF,CG,CI,CM,GA,GN,ML,MR,NE,SN,TD,TG

National: AL,AM,AT,AU,AZ,BA,BB,BG,BR,BY,CA,CH,CN,CU,CZ,DE,DK,EE,ES,FI,GB,GE,GH,HU, IL,IS,JP,KE,KG,KP,KR,KZ,LC,LK,LR,LS,LT,LU,LV,MD,MG,MK,MN,MW,MX,NO;NZ,PL,PT,RO,RU,

SD,SE,SG,SI,SK,TJ,TM,TR,TT,UA,UG,US,UZ,VN,YU

ATTENTION

The applicant should carefully check the data appearing in this Notification. In case of any discrepancy between these data and the indications in the international application, the applicant should immediately inform the International Bureau.

In addition, the applicant's attention is drawn to the information contained in the Annex, relating to:

time limits for entry into the national phase; confirmation of precautionary designations;

requirements regarding priority documents.

A copy of this Notification is being sent to the receiving Office and to the International Searching Authority.

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland

Authorized officer:

R. Raissi

Telephone No. (41-22) 730.91.11





INFORMATION ON TIME LIMITS FOR ENTERING THE NATIONAL PHASE

The applicant is reminded that the "national phase" must be entered before each of the designated Offices indicated in the Notification of Receipt of Record Copy (Form PCT/IB/301) by paying national fees and furnishing translations, as prescribed by the applicable national laws.

The time limit for performing these procedural acts is 20 MONTHS from the priority date or, for those designated States which the applicant elects in a demand for international preliminary examination or in a later election, 30 MONTHS from the priority date, provided that the election is made before the expiry of 19 months from the priority date. Some designated (or elected) Offices have fixed time limits which expire even later than 20 or 30 months from the priority date. In other Offices an extension of time or grace period, in some cases upon payment of an additional fee, is available.

In addition to these procedural acts, the applicant may also have to comply with other special requirements applicable in certain Offices. It is the applicant's responsibility to ensure that the necessary steps to enter the national phase are taken in a timely fashion. Most designated Offices do not issue reminders to applicants in connection with the entry into the national phase.

For detailed information about the procedural acts to be performed to enter the national phase before each designated Office, the applicable time limits and possible extensions of time or grace periods, and any other requirements, see the relevant Chapters of Volume II of the PCT Applicant's Guide: Information about the requirements for filing a demand for international preliminary examination is set out in Chapter IX of Volume I of the PCT Applicant's Guide.

Note that since ES is not bound by PCT Chapter II (which provides for the international preliminary examination procedure), that State cannot be elected in a demand for international preliminary examination. In the case of the designation of ES for a national patent, the applicant must thus always enter the national phase before the national Office of that State before the expiration of 20 months from the priority date. In the case of the designation of ES for a European patent, however, the 31-month time limit applies in respect of that designation if at least one other State designated for a European patent is also elected within the 19-month period.*

Note also that only an applicant who is a national or resident of a PCT Contracting State which is bound by Chapter II has the right to file a demand for international preliminary examination.

* CH and LI became bound by PCT Chapter II on 1 September 1995. GR became bound by PCT Chapter II on 7 September 1996. Therefore, CH and LI may be elected in a demand or a later election filed on or after 1 September 1995, and GR may be elected in a demand or a later election filed on or after 7 September 1996, regardless of the filing date of the international application. (See 2nd paragraph above.)

CONFIRMATION OF PRECAUTIONARY DESIGNATIONS

This notification lists only specific designations made under Rule 4.9(a) in the request. It is important to check that these designations are correct. Errors in designations can be corrected where precautionary designations have been made under Rule 4.9(b). The applicant is hereby reminded that any precautionary designations may be confirmed according to Rule 4.9(c) before the expiration of 15 months from the priority date. If it is not confirmed, it will automatically be regarded as withdrawn by the applicant. There will be no reminder and no invitation. Confirmation of a designation consists of the filing of a notice specifying the designated State concerned (with an indication of the kind of protection or treatment desired) and the payment of the designation and confirmation fees. Confirmation must reach the receiving Office within the 15-month time limit.

REQUIREMENTS REGARDING PRIORITY DOCUMENTS

For applicants who have not yet complied with the requirements regarding priority documents the following is recalled.

Where the priority of an earlier national (i.e., national or regional) application is claimed, the applicant must submit a copy of the said national application, certified by the authority with which it was filed ("the priority document") to the receiving Office (which will transmit it to the International Bureau) or directly to the International Bureau, before the expiration of 16 months from the priority date (Rule 17.1).

Where the priority document is issued by the receiving Office, the applicant may, instead of submitting the priority document, request the receiving Office to prepare and transmit the priority document to the International Bureau. Such a request must be made before the expiration of the 16-month time limit.

It is recalled that, where several priorities are claimed, the priority date to be considered for the purposes of computing the 16-month time limit is the filing date of the earliest application whose priority is claimed.

If the priority document concerned is not submitted to the International Bureau before the expiration of the 16-month time limit, or if the request to the receiving Office to transmit the priority document has not been made (and the corresponding fee, if any, paid) before the expiration of this time limit, any designated State may disregard the priority claim.



_	from the RECEIVING OFFICE					
	Fo: Mewburn Ellis York House	TOTAL SANCE		, i Nor	PCT	
	23 Kingsway London	2 2 MAY 199	7	A	PPLICATION NUMBER AND OF THE INTERNATIONAL FILING DATE	
	WC2B 6HP			Date of mailing	(PCT Rule 20.5(c)) 20/05/47.	
	Applicant's or agents's file reference SMW/BP5610415			IMI	PORTANT NOTIFICATION	
L	International application No. PCT/GB97/01250 Applicant	International filing date 08/05/1			Priority date (day/month/year) 08/05/1996	
-	University of Dundee et al Title of the invention					
L	Methods and means for inhibition					
	1. The applicant is hereby notified that the international application has been accorded the international application number and the international filing date indicated above. 2. The applicant is further notified that the record copy of the international application:					
	(with Form PCT/IB/301) of its receithe priority date, the International B	the promorme rice record CO	DV D	IOI nave been rece	ing Office and will notify the applicant sived by the expiration of 14 months from	
Na	me and mailing address of the receiving The Patent Office	ng Office	A	uthorized officer		
	Cardiff Road, Newport				J. R. LLOYD - THOMAS	
Fac	simile No. Gwent NP9 1RH		Te	elenhone Na	ROOM G.Y73 EXT. 4587: 4346.	

Form PCT/RO/105 (July 1992)

Ĵ		
	The demand must be filed directly with the comment International Preliminary Examining Author if two or more A	Authorities are competent
wi	with the one chosen by the applicant. The full me or two-letter code of that Authority may be in the applicant.	nt on the line below:
777	IDEA!	

PCT

CHAPTER II

DEMAND

under Article 31 of the Patent Cooperation Treaty:

The undersigned requests that the international application specified below be the subject of international preliminary examination according to the Patent Cooperation Treaty.

Identification of IPEA		Date of receipt of DE	MAND	
Box No. I IDENTIFICATION OF THE INTERNATIONAL APPLICA		PPLICATION	Applicant's or agent's file reference SJK/BP5610415	
International application No.	International filing of	date (day/month/year)	(Earliest Priority date (day/month/year)	
PCT/GB97/01250	08/05/1997		8 MAY 1996	
Title of invention METHODS AND I	MEANS FOR INHIBITION C	F CDK4 ACTIVITY		
Box No. II APPLICANT(S)				
Name and address: (Family name followed designation. The address)	ed by given name; for a legal enti dress must include postal code an		Telephone No.:	
CYCLACEL LIMITED MARQUIS HOUSE 67-68 JERMYN STREET	·		Facsimile No.:	
LONDON SW1Y 6NY UNITED KINGDOM	·		Teleprinter No.:	
State (i.e. country) of nationality:	GB	State (i.e. country) o	of residence: GB	
Name and address: (Family name followed	by given name; for a legal entity,	full official designation. The	address must include postal code and name of country	
BALL KATHRYN LINDSAY CRC CELL TRANSFORMATION RI DEPARTMENT OF BIOCHEMISTR' UNIVERSITY OF DUNDEE DUNDEE DD1 4HN UNITED KINGDOM		-		
State (i.e. country) of nationality:	GB	State (i.e. country)	of residence: GB	
Name and address: (Family name followed LANE DAVID PHILIP CRC CELL TRANSFORMATION REDEPARTMENT OF BIOCHEMISTRY UNIVERSITY OF DUNDEE DUNDEE DD1 4HN UNITED KINGDOM	SEARCH GROUP	I full official designation. The o	address must include postal code and name of countr	
State (i.e. country) of nationality:	GB	State (i.e. country) o	of residence: GB	

International application No. PCT/GB97/01250 BOX NO. III AGENT OR COMMON REPRESENTATIVE; OR ADDRESS FOR CORRESPONDENCE The following person agent common representative X has been appointed earlier and represents the applicant(s) also for international preliminary examination. and is hereby appointed and any earlier appointment of (an) agent(s)/common representative is hereby revoked. is hereby appointed, specifically for the procedure before the International Preliminary Examining Authority, in addition to the agent(s)/common representative appointed earlier. (Family name followed by given name; for a legal entity, full official Telephone No.: Name and address: designation. The address must include postal code and name of country.) 0171 240 4405 KIDDLE, SIMON J. and others Mewburn Ellis Facsimile No.: York House 0171 240 9339 23 Kingsway London WC2B 6HP Teleprinter No .: GB 22762 PATENT G Mark this check-box where no agent or common representative is/has been appointed and the space above is used instead to indicate a special address to which correspondence should be sent. Box No. IV STATEMENT CONCERNING AMENDMENTS The applicant wishes the International Preliminary Examining Authority* to start the international preliminary examination on the basis of the international application as originally filed. (i) (ii) to take into account the amendments under Article 34 of the description (amendments attached). the claims (amendments attached). the drawings (amendments attached). to take into account any amendments of the claims under Article 19 filed with the International Bureau (a copy is (iii) attached). to disregard any amendments of the claims made under Article 19 and to consider them as reversed. (iv) to postpone the start of the international preliminary examination until the expiration of 20 months from the priority date (v) unless that Authority receives a copy of any amendments made under Article 19 or a notice from the applicant that he does not wish to make such amendments (Rule 69.1(d)). (This check-box may be marked only where the time limit under Article 19 has not yet expired.) Where no check-box is marked, international preliminary examination will start on the basis of the international application as originally filed, or where a copy of amendments to the claims under Article 19 and/or amendments of the international application under Article 34 are received by the International Preliminary Examining Authority before it has begun to draw up a written opinion or the international preliminary examination, as so amended. Box No. V ELECTION OF STATES The applicant hereby elects all eligible States (that is, all States which have been designated and which are bound by Chapter II of the PCT) except (If the applicant does not wish to elect certain eligible States, the name(s) or country code(s) of those States must be indicated

above.)

£.		International application No.				
Box No. VI CHECK LIST		PCT/GB97/01250				
The demand is accompanied by the following documen	rs for the	For International Preliminary				
purposes of international preliminary examination:		Examining Authority use only				
1. amendments under Article 34		received not received				
description : 0	sheets					
claims : 0	sheets					
drawings : 0	sheets					
2. letter accompanying amendments under Article 34 :	sheets					
3. copy of amendments under Article 0	sheets					
4. copy of statement under Article 0	sheets					
5. other (specify) : 0	sheets					
The demand is also accompanied by the item(s) marked	l below:					
1. separate signed power of attorney	4.	fee calculation sheet				
2. copy of general power of attorney	5.	X other (specify):				
3. statement explaining lack of signature	Αſ	ODITIONAL REPRESENTATIVES				
Box No. VII SIGNATURE OF APPLICANT, A	AGENT OR	COMMON REPRESENTATIVE				
KIDDLE, SIMON J. APPOINTED AGENT						
For Internati	onal Prelimina	ary Examining Authority use only				
1. Date of actual receipt of DEMAND:						
 Adjusted date of receipt of demand due to CORRECTIONS under Rule 60.1(b): 						
The date of receipt of the demand is AFTER the expiration of 19 months from the priority date and item 4 or 5, below, does not apply. The applicant has been informed accordingly.						
4. The date of receipt of the demand is WITHIN the period of 19 months from the priority date as extended by virtue of Rule 80.5						
5. Although the date of receipt of the demand is after the expiration of 19 months from the priority date, the delay in arrival is EXCUSED pursuant to Rule 82.						
For I	nternational B	ureau use only				
Demand received from IPEA on:						

REQUEST

WALTON, SEAN M. and others MEWBURN ELLIS YORK HOUSE 23 KINGSWAY

NDON WC2B 6HP

D.C.W.	For receiving Office use only				
PCT					
REQUEST	International Application No.				
The undersigned requests that the present international application be processed	International Filing Date				
according to the Patent Cooperation Treaty	Name of receiving Office and "PCT" International Application"				
	Applicant's or agent's file reference SMW/BP5610415 (if desired) (12 characters maximum)				
Box No. I TITLE OF INVENTION METHODS AND	MEANS FOR INHIBITION OF CDK4 ACTIVITY				
Box No. II APPLICANT					
Name and address: (Family name followed by given name; for a legal designation. The address must include postal cod	l entity, full official de and name of country.) This person is also inventor.				
UNIVERSITY OF DUNDEE Dundee DD1 4HN	Telephone No.				
UNITED KINGDOM	Facsimile No.				
	Teleprinter No.				
	State (i.e. country) of residence: GB				
This person is applicant for the purposes of: all designated States X all designated States	the United States of the States indicated in the of America only Supplemental Box				
Box No. III FURTHER APPLICANT(S) AND/OR (FU	RTHER) INVENTOR(S)				
Name and address: (Family name followed by given name; for a legal designation. The address must include postal code	e and name of country.) This person is:				
BALL Kathryn Lindsay CRC Cell Transformation Research Group Department of Biochemistry	applicant only				
University of Dundee Dundee DD1 4HN UNITED KINGDOM	X applicant and inventor				
	inventor only (if this check-box is marked, do not fill in below.)				
State (i.e. country) of nationality: GB	State (i.e. country) of residence: GB				
This person is applicant for the purposes of: all all designated State United States of A	es except the X the United States the States indicated in the Supplemental Box				
X Further applicants and/or (further) inventors are indicated on a continuation sheet.					
Box No. IV AGENT OR COMMON REPRESENTATIVE; OR ADDRESS FOR CORRESPONDENCE					
The person identified below is hereby/has been appointed to act on behalf of the pplicant(s) before the competent International Authorities as:					
Name and address: (Family name followed by given name; for a legal entity, The address must include postal code and name of country	full official designation. Telephone No. 0171 240 4405				

Mark this check-box where no agent or common representative is/has been appointed and the space above is used instead to indicate a special address to which correspondence should be sent. Form PCT/RO/101 (first sheet) (5 July 1994: reprint July 1996)

Facsimile No. 0171 240 9339

Teleprinter No. 22762 PATENT G

Continuation of Box No. III FURTHER APPLICANTS AND/OR (FURTHER) INVENTORS					
If none of the following sub-boxes is used, this sheet is not to be included in the request.					
Name and address: (Family name followed by given name; for a lego designation. The address must include postal co	nl entiry, full official de and name of country.)	This person is:			
LANE David Philip CRC Cell Transformation Research Group Department of Biochemistry	applicant only				
CRC Cell Transformation Research Group Department of Biochemistry University of Dundee Dundee DD1 4HN UNITED KINGDOM		X applicant and inventor			
		inventor only (if this check-box is marked, do not fill in below.)			
State (i.e. country) of nationality: GB	State (i.e. country) of resid	ence: GB			
This person is applicant for all designated the purposes of: all designated States United States of		United States America only the States indicated in the Supplemental Box			
Name and address: (Family name followed by given name; for a legal designation. The address must include postal co	al entity, full official de and name of country.)	This person is:			
		applicant only			
		applicant and inventor			
		inventor only (if this check-box is marked, do not fill in below.)			
State (i.e. country) of nationality:	State (i.e. country) of resid	ence:			
This person is applicant for all designated the purposes of: all designated States United States o	• 1 1	United States the States indicated in the Supplemental Box			
Name and address: (Family name followed by given name; for a legal entity, full official designation. The address must include postal code and name of country.)					
		applicant only			
		applicant and inventor			
		inventor only (if this check-box is marked, do not fill in below.)			
State (i.e. country) of nationality:	State (i.e. country) of resid	lence:			
This person is applicant for all designated the purposes of: all designated States United States of	• • •	United States America only the States indicated in the Supplemental Box			
Name and address: (Family name followed by given name; for a legal designation. The address must include postal co		This person is:			
		applicant only			
		applicant and inventor			
		inventor only (if this check-box is marked, do not fill in below.)			
State (i.e. country) of nationality: State (i.e. country) of residence:					
This person is applicant for all designated the purposes of: all designated States except the United States of America only the United States indicated in the Supplemental Box					
Further applicants and/or (further) inventors are indicated on another continuation sheet.					

Box	No. V	DESIGNATION OF STATES		_
The Reg	'owin	g designations are hereby made under Rule 4.9(a) (mark the atent	e appl	icable check-boxes; at least one must be marked):
X		ARIPO Patent: KE Kenya, LS Lesotho, MW Malawi, SI is a Contracting State of the Harare Protocol and of the PC	CT	
X	EA	Eurasian Patent: AM Armenia, AZ Azerbaijan, B of Moldova, RU Russian Federation, TJ Tajikistan, Contracting State of the Eurasian Patent Convention	TM	larus, KG Kyrgyzstan, KZ Kazakstan, MD Republic Turkemenistan, and any other State which is a f the PCT
X	EP	European Patent: AT Austria, BE Belgium, CH and Denmark, ES Spain, FI Finland, FR France, GB Un Luxembourg, MC Monaco, NL Netherlands, PT Por Contracting State of the European Patent Convention	nited I rtugal	Kingdom, GR Greece, IE Ireland, IT Italy, LU, SE Sweden, and any other State which is a
X	OA	OAPI Patent: BF Burkina Faso, BJ Benin, CF Cen Cameroon, GA Gabon, GN Guinea, ML Mali, MR and any other State which a member State of OAPI a protection or treatment desired, specify on dotted lin	Maur and a	itania, NE Niger, SN Senegal, TD Chad, TG Togo, Contracting State of the PCT (if other kind of
Nat	ional P	atent (if other kind of protection desired, specify on a	lotted	line):
		Albania	X	LU Luxembourg
	AM	Armenia	X	LV Latvia
X	AT		X	MD Republic of Moldova
团	ΑU	Australia	X	MG Madagascar
团	AZ	Azerbaijan	X	MK The former Yugoslav Republic of Macedonia
团	BA			MN Mongolia
\square	BB	Barbados	X	MW Malawi
团	BG	Bulgaria	X	MX Mexico
区	BR	Brazil	X	NO Norway
团	BY	Belarus	XXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXX	NZ New Zealand
X	CA	Canada	冈	PL Poland
X	CH	and LI Switzerland and Liechtenstein	团	PT Portugal
X	CN	China	X	RO Romania
対	CU	Cuba	X	RU Russian Federation
X	\mathbf{CZ}	Czech Republic	冈	SD Sudan
冈	DE	Germany	X	SE Sweden
X	DK	Denmark	X	SG Singapore
X	EE	Estonia	X	SI Slovenia
X	ES	Spain	$\overline{\mathbf{X}}$	SK Slovakia
X	FI	Finland	X X X	TJ Tajikistan
X	GB	United Kingdom	X	TM Turkmenistan
X	GE	Georgia	$\overline{\mathbf{X}}$	TR Turkey
X	HU	Hungary	X	TT Trinidad and Tobago
X	TL.	Israel	X X X X	UA Ukraine
X	IS	Iceland ,	X	UG Uganda
X	JР	Japan	X	US United States of America
X	KE	Kenya	X	UZ Uzbekistan
X	KG	Kyrgyzstan	X	VN Viet Nam
X	KP	Democratic People's Republic of Korea		
X	KR	Republic of Korea		k-boxes reserved for designating States (for the
X	KZ	Kazakstan		oses of a national patent) which have become party to
	LC	St Lucia	ine P	PCT after issuance of this sheet:
X	LK	Sri Lanka	\square	YU Yugoslavia
X	LR	Liberia	X	GH Ghana
X		Lesotho	M	All and any further PCT States not listed
X		Lithuania		on this form
In	addition	to the designations made above, the applicant also makes	unde	r Kule 4.9(b) all designations which would be permitted

under the PCT except the designation(s) of
The applicant declares that those additional designations are subject to confirmation and that any designation which is not confirmed
before the expiration of 15 months from the priority date is to be regarded as withdrawn by the applicant at the expiration of that time
limit. (Confirmation of a designation consists of the filing of a notice specifying that designation and the payment of the designation and
confirmation fees. Confirmation must reach the receiving Office within the 15-month time limit.)

If the Supplemental Box is not used, this sheet need not be included in the request.

l' his box in the following cases:

I. If, in any of the Boxes, the space is insufficient to furnish all the information:

in particular:

- (i) if more than two persons are involved as applicants and/or inventors and no "continuation sheet" is available:
- (ii) if, in Box No. II or in any of the sub-boxes of Box No. III, the indication "the States indicated in the Supplemental Box" is checked:
- (iii) if, in Box No. II or in any of the sub-boxes of Box No. III, the inventor or the inventor/applicant is not inventor for the purposes of all designated States or for the purposes of the United States of America:
- (iv) if, in addition to the agent(s) indicated in Box No. IV, there are further agents:
- (v) if, in Box No. V, the name of any State (or OAPI) is accompanied by the indication "patent of addition," or "certificate of addition," or if, in Box No. V, the name of the United States of America is accompanied by an indication "Continuation" or "Continuation-in-part":
- (vi) if there are more than three earlier applications whose priority is claimed:
- 2. If the applicant claims, in respect of any designated Office, the benefits of provisions of the national law concerning non-prejudicial disclosures or exceptions to lack of novelty:

In such case, write "Continuation of Box No. ..." (indicate the number of the Box) and furnish the information in the same manner as required according to the captions of the Box in which the space was insufficient;

in such case, write "Continuation of Box III" and indicate for each additional person the same type of information as required in Box No. III;

in such cases write "Continuation of Box No. II" or "Continuation of Box No. III" or "Continuation of Boxes No. II and No. III" (as the case may be), indicate the name of the applicant(s) involved and next to (each) such name, the State(s) (and/or, where applicable, ARIPO, Eurasian, European or OAPI patent) for the purposes of which the named person is applicant;

in such case, write "Continuation of Box No. II" or "Continuation of Box No. III" or "Continuation of Boxes No. II and No. III" (as the case may be), indicate the name of the inventor(s) and, next to (each) such name, the State(s) (and/or, where applicable, ARIPO, Eurasian, European or OAPI patent) for the purposes of which the named person is inventor;

in such case, write "Continuation of Box No. IV" and indicate for each further agent the same type of information as required in Box No. IV;

in such case, write "Continuation of Box No. V" and the name of each State involved (or OAPI), and after the name of each such State (or OAPI), the number of the parent title or parent application and the date of grant of the parent title or filing of the parent application;

in such case, write "Continuation of Box No. VI" and indicate for each additional earlier application the same type of information as required in Box No. VI.

in such case, write "Statement Concerning Non-Prejudicial Disclosures or Exceptions to Lack of Novelty" and furnish that statement below.

Continuation of Box IV

ARMITAGE, IAN M.

BRASNETT, ADRIAN H.

BREWSTER, ANDREA R.

CALDERBANK, T. ROGER

COLEIRO, RAYMOND

FORD, MICHAEL F.

GURA, H. ALAN

HACKNEY, NIGEL J.

HAMILTON, ALISTAIR

HARRISON, DAVID C.

KIDDLE, SIMON J.

LINN, S. JONATHAN

LYONS, JUNE, M.

NICHOLLS, KATHRYN M.

O'BRIEN, CAROLINE J.

PAGET, HUGH C.E.

SANDERSON, MICHAEL J.

STONER, G. PATRICK

STUART, IAN

WALTON, SEÁN M.

Box No. VI PRIORITY	CLAIM		Further priority claims	indicated in the Supplemental Box
The priority of the following ear	lier application(s) is her	reby claimed:		
Country (in which, or for which the application was filed)	Filing Date (day/month/ye		Application No.	Office of filing (only for regional or international application)
item (1) GB	8 May 199 (08/05/96		9609521.1	
item (2) GB	9 October 19 (09/10/9		9621314.5	
item (3)		·		
The receiving Office is her Bureau a certified copy of	is the receiving Office of the communication is the earlier application (s	(a fee may be re and transn s) identified a	e required): nit to the International bove as item(s): (1) & (2)	Office which for the purposes of the
Box No. VII INTERNATIO	ONAL SEARCHING A	AUTHORIT	Y	
heen carried out or requested an	sternational search, ind earch (international, int ed the Authority is now	icate the Auth ternational-ty requested to	hority chosen; the two-letter pe or other) by the Internati base the international searc	code may be used): ISA/ onal Searching Authority has already h. to the extent possible, on the results of
that earlier search. Identify such Country (or regional Office):	h search or request eith Day (day/month/)	er by referen	ce to the relevant applicatio Number:	n (or the translation thereof) or by
Box No. VIII CHECK LIST	r			
This international application couthe following number of sheets:	ntains	This interna	ational application is accomp	panied by the item(s) marked below:
1. request : 5	sheets		parate signed wer of attorney	5. K fee calculation sheet
2. description : 78	sheets	2. Cor	oy of general	6. separate indications concerning
3. claims : 11 4. abstract : 1	sheets sheets	1	wer of attorney	deposited microorganisms 7. nucleotide and/or amino acid
5. drawings : 10	sheets		tement explaining lack of nature	7. nucleotide and/or amino acid sequence listing (diskette)
Total : 105	sheets		ority document(s) entified in Box No. VI as m(s):	8 other (specify): Form 23/77 (x2)
Figure No. 0 of the	ne drawings (if any) sho	ould accompa	any the abstract when it is p	ublished.
Box No. IX SIG	NATURE OF APPLIC	CANT OR A	GENT	
Next to each signature indicate the na	me of the person signing ar	nd the capacity	in which the person signs (if such	h capacity is not obvious from reading the reques
	•			
BRASNETT, ADRIAN H.				
		For receivin	g Office use only	
Date of actual receipt of the p international application:	urported	TOT TOOCHVIII	, S	2. Drawings:
Corrected date of actual receiting received papers or draw the purported international appropriate received papers.	vings completing			received:
Date of timely receipt of the corrections under PCT Artic	·			not received:
International Searching Auth the applicant: ISA/			nittal of search copy delayed earch fee is paid	d
Date of receipt of the record co		r Internations	al Bureau use only	

٦,	This sheet is no	ot parragand does not cou	int as a she			
	PC			For receiving ff	ice use only	<i>,</i>
	FEE CALCULAT Annex to the		Inte	rnational application No). 	
			·			
	licant's or agent's reference	SMW/BP5610415	Dat	e stamp of the receiving	g Office	
App	licant UNIVE	RSITY OF DUNDEE				
CA	LCULATION OF PRESCRIBED	FEES				
	TRANSMITTAL FEE			1 £55	T	
2.	SEARCH FEE		`	£1003	S	
	International search to be carried (If two or more International Searchiname of the Authority which is chosen	ng Authorities are competent in re	elation to the ir	nternational application, in	dicate the	
3.	INTERNATIONAL FEE					
	Basic Fee The international application cont	ains 105 sheets.				
	first 30 sheets			Ь		
					İ	
	75 x £6 remaining sheets additional a			b		
	Add amounts entered at b ₁ and b ₂	and enter total at B	£778	В		
	Designation Fees					
	The international application cont	ains 71 designations.				
	xx	£80 =	£880	D		
	number of designation fees amo payable (maximum 11)	ount of designation fee				
	Add amounts entered at B and D (Applicants from certain States are en international fee. Where the applicant	utitled to a reduction of 75% of th	he titled. the	£1658	I	
	total to be entered at I is 25% of the	sum of the amounts entered at B	and D.)	• • • •		
	FEE FOR PRIORITY DOCUME			£44	P	
3	Add amounts entered at T, S, I a	TOTAL FEES and P, and enter total in the T		£2760		9.7
				TOTAL		
				151		
Г	The designation fees are not pa	d at this time.				
MO	DDE OF PAYMENT					4
	authorization to charge deposit account (see below)	bank draft		upons		
X	cheque	cash	oth	ner (specify)		·
	postal money order	revenue stamps				
DE	POSIT ACCOUNT AUTHORIZ	ATION (this mode of payme	nt may not be	e available at all receiv	ing Offices)	·
Th	e RO/ is hereby auth	orized to charge the total fee	indicated ab	ove to my deposit accor	unt.	
	deposit accou					
		norized to charge the fee for IPO to my deposit account.	preparation a	nd transmittal of the pri	ority docur	nent to the International

Signature

From the INTERNATIONAL SEARCHING AUTHORITY	PCT
	NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION PCT Rule 44.1) Face of mailing (day month year) 1 7 00 07
	(day/month/year) 17,09,07
Applicant's or agent's file reference SMW/BP5610415 International application No.	FOR FURTHER ACTION See paragraphs 1 and 4 below International filing date
PCT/GB 97/01250	(day month year) 08/05/1997
Applicant	
CYCLACEL LIMITED et al.	
1. X The applicant is hereby notified that the International Search Filing of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claims When? The time limit for filing such amendments is normal International Search Report; however, for more de Where? Directly to the International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	s of the International Application (see Rule 46):
Fascimile No.: (41-22) 740.14.35	
For more detailed instructions, see the notes on the accompand. 2. The applicant is hereby notified that no International Search Article 17(2)(a) to that effect is transmitted herewith.	·
3. With regard to the protest against payment of (an) additiona	l fee(s) under Rule 40.2, the applicant is notified that:
the state of the s	n transmitted to the International Bureau together with the rotest and the decision thereon to the designated Offices.
no decision has been made yet on the protest; the appl	licant will be notified as soon as a decision is made.
4. Further action(s): The applicant is reminded of the following: Shortly after 18 months from the priority date, the international a If the applicant wishes to avoid or postpone publication, a noti- priority claim, must reach the International Bureau as provided completion of the technical preparations for international publications.	in Rules 90bis.1 and 90bis.3, respectively, before the
Within 19 months from the priority date, a demand for internatio wishes to postpone the entry into the national phase until 30 m	nal preliminary examination must be filed if the applicant nonths from the priority date (in some Offices even later).
Within 20 months from the priority date, the applicant must perform the before all designated Offices which have not been elected in the priority date or could not be elected because they are not boun	delitation of the a facer electron manager
Name and mailing address of the International Searching Authority	Authorized officer
European Patent Office, P.B. 5818 Patentlaan 2 NL-2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Alfredo Prein

NOTES TO FORM PCT/ISA/220

These Notes are intended to give the basic instructions concerning the filing of amendments under article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions respectively.

INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international pbulication. Furthermore, it should be emphasized that provisional protection is available in some States only.

What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When?

Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been/is filed, see below.

How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

What documents must/may accompany the amendments?

Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

NOTES TO FORM PCT/ISA/220 (continued)

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled;
- (iii) the claim is new;
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- [Where originally there were 48 claims and after amendment of some claims there are 51]:
 "Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers;
 claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- [Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- 3. [Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims]:
 "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or
 "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- 4. [Where various kinds of amendments are made]: "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claim 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

"Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

It must be in the language in which the international appplication is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended. It must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

Consequence if a demand for international preliminary examination has already been filed

If, at the time of filing any amendments under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the same time of filing the amendments with the International Bureau, also file a copy of such amendments with the International Preliminary Examining Authority (see Rule 62.2(a), first sentence).

Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, where upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's

PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference	FOR FURTHER see Notification o (Form PCT/ISA/	f Transmittal of International Search Report 220) as well as, where applicable, item 5 below.
International application No.	International filing date(day month year)	(Earliest) Priority Date (day/month/year)
PCT/GB 97/01250	08/05/1997	08/05/1996
Applicant		
CYCLACEL LIMITED et al.		
This International Search Report has bee according to Article 18. A copy is being to	n prepared by this International Searching Autransmitted to the International Bureau.	thority and is transmitted to the applicant
This International Search Report consists [X] It is also accompanied by a cop	s of a total of 5 sheets. by of each prior art document cited in this repo	rt.
1. X Certain claims were found unsea	archable (see Box I).	
2. Unity of invention is lacking (see	e Box II).	
3. X The international application of international search was carried	ontains disclosure of a nucleotide and/or amino I out on the basis of the sequence listing	acid sequence listing and the
file	d with the international application.	
X fur	nished by the applicant separately from the int	
	but not accompanied by a statement to the matter going beyond the disclosure in the	e international application as filed.
Tra	anscribed by this Authority	
4 Wish second to the site. W the	text is approved as submitted by the applicant	L
,	text has been established by this Authority to	
5. With regard to the abstract,	and a submitted by the employer	
	e text is approved as submitted by the applican text has been established, according to Rule 3	8.2(b), by this Authority as it appears in
⊢ Bo	ox III. The applicant may, within one month frarch Report, submit comments to this Authori	om the date of mailing of this international
6. The figure of the drawings to be put		
	suggested by the applicant.	None of the figures.
\ \	cause the applicant failed to suggest a figure. cause this figure better characterizes the invent	ion.
	cause and infaire octors offer percentage and investor	

ernational application No.

INTERNATIONAL SEARCH REPORT

PCT/GB 97/01250

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: 31-35, 38-41 because they relate to subject matter not required to be searched by this Authority, namely: Remark: Although claim(s) 31-35, 38-41 is(are) directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
1. As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

A. CLASSIFICATION OF SUBJECT MATTER IPC 6 C07K14/47 A61K38/17 G01N33/68

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC 6 C07 K A61 K G01 N

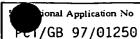
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUM	MENTS CONSIDERED TO BE RELEVANT	
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Х	MOL.CELL.BIOL., vol. 16, no. 4, April 1996, pages 1786-1793, XP002039986 LIN E.A.: "Anylysis of wild-type and mutant p21(waf1) gene activities"	1-9,29, 30
Y	The whole document; see especially page1787, column 1	1-3, 10-22, 29-35, 38-41
Y	NATURE, vol. 375, 11 May 1995, LONDON GB, pages 159-161, XP002039811 LUO E.A.: "Cell-cycle inhibition by independent CDK and PCNA binding domains in p21(cip1)" cited in the application See especially p.159, column 2	1-3, 10-22, 29-35, 38-41
	-/	

X Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
* Special categories of cited documents: A' document defining the general state of the art which is not considered to be of particular relevance E' earlier document but published on or after the international filing date	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed	involve an inventive step when the document is taken alone 'Y' document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such docu- ments, such combination being obvious to a person skilled in the art. '&' document member of the same patent family
Date of the actual completion of the international search 8 September 1997	Date of mailing of the international search report 1 7, 09, 97
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer Groenendijk, M

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	tion) DOCUMENTS CONSIDERED TO BE RELEVANT	In the second second
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	ONCOGENE, vol. 12, no. 3, 1 February 1996, pages 595-607, XP002039814 CHEN E.A.: "Characterization of p21(cip1/waf1) peptide domains required for cyclin E/Cdk2 and PCNA interaction" cited in the application The whole document; see especially p.599	1-3, 10-22, 29-35, 38-41
Y	EMBO JOURNAL, vol. 14, no. 3, 1995, pages 555-563, XP002039812 NAKANISHI E.A.: "Identification of the active region of the DNA synthesis inhibitory gene P21" cited in the application see the whole document	1-3, 10-22, 29-35, 38-41
Y	ONCOGENE, vol. 10, no. 12, 15 June 1995, pages 2281-2287, XP002039813 GOUBIN E.A.: "Identification of binding domains on the p21(cip1) cyclin-dependent kinase inhibitor" cited in the application see the whole document	1-3, 10-22, 29-35, 38-41
P,X	CURRENT BIOLOGY, vol. 7, no. 1, 1 January 1997, pages 71-80, XP002039815 BALL E.A.: "Cell-cycle arrest and inhibition of Cdk4 activity by small peptides based on the carboxy-terminal domain of p21waf1" see the whole document	1-41
P,X	MOL.CELL.BIOL., vol. 16, no. 9, September 1996, pages 4673-4682, XP002039987 CHEN E.A.: "Cyclin-binding motifs are essential for the function of p21(cip1)" see the whole document	. 1-41
A	CURRENT BIOLOGY, vol. 5, 1 January 1995, pages 275-282, XP000607149 WARBRICK E ET AL: "A SMALL PEPTIDE INHIBITOR OF DNA REPLICATION DEFINES THE SITE OF INTERACTION BETWEEN THE CYCLIN-DEPENDENT KINASE INHIBITOR P21WAF1 AND PROLIFERATING CELL NUCLEAR ANTIGEN" cited in the application see the whole document	1-41
	-/	

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ategory *	tion) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
	NISHINIHON JOURNAL OF UROLOGY / NISHI NIHON HIYOKIKA, vol. 5, no. 2, 1 January 1995, pages 127-156, XP000572011 MACLACHLAN T K ET AL: "CYCLINS, CYCLIN-DEPENDENT KINASES AND CDK INHIBITORS: IMPLICATIONS IN CELL CYCLE CONTROL AND CANCER"	1-41
	FR 2 662 698 A (CENTRE NAT RECH SCIENT) 6 December 1991 see the whole document	35

INTERNAL SEARCH REPORT

on on patent family members

1	In onal Application No
	PCT/GB 97/01250

Patent document cited in search report	Publication	Patent family	Publication
	date	member(s)	date
FR 2662698 A	06-12-91	AT 143691 T DE 69122472 D DE 69122472 T EP 0485578 A WO 9118981 A JP 5502885 T	15-10-96 07-11-96 17-04-97 20-05-92 12-12-91 20-05-93